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Abstract: BACKGROUND: A history of illicit injecting drug use makes indication of total hip arthroplasty (THA) in patients with end stage hip osteoarthritis difficult, as the risk of infection with colonized strains is multiplied if the patient continues to inject or inhale illicit drugs. METHODS: A retrospective survivorship analysis of a consecutive series of 27 THA in patients with a history of illicit drug use was performed. Follow-up evaluation consisted of (1) a WOMAC score, (2) a standardized interview including queries on drug habits and eventual additional medico-surgical treatments of the affected hip, (3) a clinical examination in order to complete a Harris Hip Score, (4) radiological examination and (5) blood tests (blood sedimentation rates and C-reactive protein). Defined endpoints were death, implant revised or awaiting revision for deep infection or any other reason and lost to follow-up or follow-up after at least 2 years. RESULTS: Overall, 5- and 10-year implant survival rates with failure for any reason were 61 % (CI: 41;81) and 52.3 % (CI: 29;76) and for septic reasons 70.6 % (CI: 52;89) and 60.5 % (CI: 36;85), respectively. Even if at the time of THA all patients and respective health care professionals confirmed abstinence of illicit injecting drug use, five patients reported occasional use. Declared abstinence of less than 1 year before THA was associated with higher recurrence rates ($p = 0.001$) and both with higher septic failure rates ($p = 0.023$, $p = 0.061$). Positive serology for human deficiency virus did not increase implant failure rates. CONCLUSION: We use this unacceptable high failure rate as evidence when counseling patients and their health care professionals about the appropriate treatment of osteoarthritis in patients with a history of illicit drug use. Furthermore, we support the request of hair analysis for drugs documenting abstinence of at least 1 year before indicating THA.

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Abstract

Background A history of illicit injecting drug use makes indication of total hip arthroplasty (THA) in patients with end stage hip osteoarthritis difficult, as the risk of infection with colonized strains is multiplied if the patient continues to inject or inhale illicit drugs.

Methods A retrospective survivorship analysis of a consecutive series of 27 THA in patients with a history of illicit drug use was performed. Follow-up evaluation consisted of (1) a WOMAC score, (2) a standardized interview including queries on drug habits and eventual additional medico-surgical treatments of the affected hip, (3) a clinical examination in order to complete a Harris Hip Score, (4) radiological examination and (5) blood tests (blood sedimentation rates and C-reactive protein). Defined endpoints were death, implant revised or awaiting revision for deep infection or any other reason and lost to follow-up or follow-up after at least 2 years.

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Conclusion We use this unacceptable high failure rate as evidence when counseling patients and their health care professionals about the appropriate treatment of osteoarthritis in patients with a history of illicit drug use. Furthermore, we support the request of hair analysis for drugs documenting abstinence of at least 1 year before indicating THA.

Keywords Hip arthroplasty · Survivorship analysis · Illicit drug abuse

Introduction

Illicit injecting drug use has a long history, but it was in the early 1980s, with the rapid growth of intravenous heroine use and the spread of the human immunodeficiency virus (HIV), that this behavior gained prominence as a core element of Europe's and the world's drug problem. Today the estimated prevalence of injecting drug users is about 2.5 per 1,000 individuals aged 15–64 years, which corresponds to a total number of up to one million active injecting drug users in the 27 Member States of the European Union [1].

In developed countries, illicit injecting drug users have an increased risk of premature death from drug overdose, violence, suicide, alcohol-related causes and AIDS. Furthermore, the prevalence of infection with hepatitis B, C and human immunodeficiency virus is estimated to be greater than 60 % [2] and 12 % [3], respectively. In HIV patients undergoing antiviral therapy, the risk of osteonecrosis of the femoral head (ONFH) is increased [4–6], with a prevalence of approximately 0.74 % [4]. ONFH, together with inflammatory arthropathies due to repetitive bacteremia [7] and posttraumatic conditions are the main reasons for secondary hip osteoarthritis in illicit injecting drug users.

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The treatment strategy for illicit injecting drug users primarily aims to reduce drug injection-related harm and gradual reintegration of patients into normal working life and assimilation into the social community. Secondary hip osteoarthritis may sometimes compromise return to work and thus social reintegration. In such conditions orthopedic surgeons may be approached for adequate treatment of hip osteoarthritis. When total hip arthroplasty (THA) is anticipated, the risk of septic complications is a major concern, since the risk to cause and spread infection with colonized strains is known to be multiplied by injection or inhalation of drugs [8, 9]. On the other hand, alternatives such as hip arthrodesis or resection arthroplasty may hardly be perceived to give the highest attainable standard of care by patients and their supporting health care professionals, especially if promising efforts to stop illicit drug use have been made.

During the last decade it has been the practice of our institution to accept patients with a history of illicit injecting drug use for THA if they were in medical and social care by professionals specialized in drug-related problems and if abstinence from illicit drug use was declared by patients and confirmed by their respective healthcare professionals. It has been our impression that the septic complication rates were higher than expected from a normal population. Because of absence of any data on this issue, a retrospective review of all patients with a history of illicit drug use receiving THA in our department during the last decade was undertaken.

Materials and methods

Our prospectively collected database on THA was retrospectively analyzed for patients with a history of illicit injecting drug use undergoing THA between January 1999 and December 2009. From 2,625 primary THA performed during this time period, a consecutive series of 24 patients (27 THA), 17 male and 7 female, with an average of 43 years (range 28–55) was identified.

The study was approved by our institutional Human Subject Review Board. Written consent was obtained from all patients willing to attend a follow up visit. For those denying, based on a general permit issued by the responsible state agency, our institutional review board allows retrospective analysis of patient data relating to standard diagnostic or therapeutic procedures without individual informed consent. There was no external funding for the purpose of this study.

Treatment protocol

Before considering patients for THA, abstinence from illicit injecting drug use had to be declared by patients and

confirmed by respective health care professionals and, in case of positive serology for hepatitis C virus (HCV) and/or HIV, patients had to be followed by a healthcare institution specialized in infectious disease. Additionally, blood sedimentation rates (BSR) less than 35 mm/h and C-reactive protein (CRP) levels less than 10 mg/L on preoperative blood screenings were mandatory.

Surgeries were performed under a laminar air flow system. Depending on the time period and underlying pathology, either a mini-incision direct anterior approach in the supine position (10 THA), or a transgluteal (14 THA) or transtrochanteric (3 THA) approach in the lateral position was used. Three doses of cefuroxime 1.5 g in an 8-h interval starting 30 min before incision and enoxaparin sodium on a daily dose for 6 weeks were given to all patients. Depending on the time period, non-cemented (14 THA) or hybrid (non-cemented cup and cemented stem) (13 THA) implants were used. One closed suction drain lying under the fascia was used and removed after 48 h in all cases.

Follow-up evaluation

All patients were invited by mail and asked for informed consent to participate in our review. Follow-up evaluation consisted of (1) a WOMAC score, (2) a standardized interview including queries on drug habits and eventual additional medico-surgical treatments of the affected hip, (3) physical examination looking for clinical signs of infection (open sinus, swelling, tenderness, warmth) in the hip region and in order to complete a Harris Hip Score (HHS), (4) radiological examination including an antero-posterior and cross-table lateral view, which were compared to the first postoperative X-rays and screened for radiolucent lines [10, 11], periosteal new bone formation and endosteal scalloping and (5) blood tests (BSR and CRP).

For patients not responding to our mail, our standardized interview was conducted by phone. For all patients a chart review recording all perioperative and postoperative complications, and in patients lost, an analysis of the latest available X-rays was performed. Defined endpoints were death, implant revised or awaiting revision for septic or any reason, lost to follow-up or follow-up after at least 2 years.

Septic failure was diagnosed if an open sinus, an abscess formation, frank pus during surgery or positive intraoperative culture from a subsequent surgical intervention were recognized. Additionally, deep sepsis was suspected if at least three of the following criteria were present: (1) erythematous, swollen, fluctuating and tender surgical site, (2) fever, chills or sweating, (3) old closed sinus, (4) elevated BSR >35 mm/h and CRP >10 mg/L in the absence of recent surgery, neoplasia, remote infection or other inflammatory conditions, (5) periosteal new bone

formation with or without endosteal scalloping or early rapidly progressive radiolucent lines and (6) positive joint aspiration.

Statistical analysis

Kaplan–Meier survivorship curves were compiled with implant failure for septic and for any reason as endpoints, respectively. For comparisons of categorical data between groups a two sided Fisher Exact Probability Test was used. All statistical analyses were performed using SPSS 17.0 for Windows. The significance level was set at $p < 0.05$.

Results

Indications for THA are summarized in Table 1. Seven patients had previous surgery on the affected hip a mean of 6.6 years (range 0.3–31; SD 12.1) before the index THA. Among these were two patients who had resection arthroplasty because of septic arthritis before. The course of all patients is summarized in Table 2.

Six patients (8 THA's) were lost to follow-up or deceased

Two of them (3 THA's) left for a foreign country and could not be traced. One, left after septic implant revision (*Escherichia coli*) 2 months after the index procedure, was included in survivorship analysis. The other four patients (5 THA's) died after a mean postoperative time of 24 months (range 6–73, SD 33) unrelated to the surgery. Three had an uneventful postoperative course. And one had a septic revision 5 weeks after the index procedure (coagulase negative *Staphylococcus*).

Four patients with telephone interview and chart review

Four refused a follow-up visit but telephone interview and chart review revealed an uneventful course in two (24 and 98 months after THA) and septic revision in two, 24 (*Staphylococcus aureus*) and 69 months (coagulase negative *Staphylococcus*) after the index procedure.

Table 1 Indications for THA

Primary OA	7
Secondary OA	
ONFH	10
Postinfectious OA	2
Posttraumatic OA	6
Childhood's disease	2

Fourteen patients available for follow-up visit

After an average time of 69 (range 24–123; SD 38) months, nine patients (9 THA's) had an uneventful course, without any clinical signs of infection and perfect radiological appearance of the THA. CRP levels and ESR were normal in seven of them. Two had elevated values for CRP (25 and 19) and BSR (38 and 29), one after a recent pneumonia, and the other without detectable reason. There overall mean HHS and WOMAC were 91 (range 71–100) and 0.8 (range 0–2), respectively.

Two patients sustained a traumatic periprosthetic fracture. One Vancouver type B2 fracture was recognized 11 months after the index procedure when loosening of the implant together with a non-union was observed. Intraoperative tissue specimens sent for microbiological examination were negative. The other (Vancouver type B2) fracture was due to a road traffic accident 6 months after index procedure. This was treated with a long revision stem in a foreign country; however this was complicated with a postoperative deep infection (*Staphylococcus aureus*). Both showed normal values for CRP and ESR at follow-up examination. HHS (99 and 100 points) and WOMAC scores (0 and 0 points) were excellent.

One patient was revised due to periprosthetic infection (coagulase negative *Staphylococci*) 1 month after index procedure. Treatment consisted of a one-stage implant revision and ciprofloxacin and rifampicin for 3 months. At follow-up, 27 months after the index procedure, no local signs of infections were encountered and X-rays were excellent. CRP measured 16 and ESR 25 mm/h. HHS was 89 and the WOMAC 0 points.

One patient with bilateral THA showed 30 months after index surgery elevated CRP (26) and BSR (52) levels as well as implant loosening with gross metaphyseal and acetabular osteolysis on both sides (Fig. 1), but denied further investigation such as hip aspiration for bacteriological tests. However, due to clinical and radiological findings, both hips were considered as septic failures.

One patient underwent surgical implant revision due to deep infection with a total of three two-stage exchanges because of recurrent infections, first with *S. aureus* 6 months after the index procedure, second with *Streptococcus mitis* and third with *Streptococcus sanguis* and *Candida albicans*. A resection arthroplasty was finally performed.

Overall, from 25 THA (in 23 patients) followed until implant failure, death or for a minimum of 24 months, 8 were revised and 2 are awaiting revision. As a cause of failure, deep sepsis was detected in 8 cases. Five- and 10-year implant survival rates with failure for any reason were 61 % (CI: 41;81) and 52.3 % (CI: 29;76) and for septic reasons 70.6 % (CI: 52;89) and 60.5 % (CI: 36;85), respectively. Figure 2 shows Kaplan–Meier survivorship analysis.

Table 2 Main parameters of all patients in the order of their appearance in the result section

Patient	THA	Gender	Age	Positive serology (years)	Side	IIDU abstinence	IIDU recurrence	MCHPP	Smoking	Alcohol	Postoperative course	Endpoint	Time (months)	CRP	BSR	HHS	WOMAC
1	1	M	34		L	>1	No	No	No	No	Uneventful	Lost	1				Excluded
2	2				R						Uneventful						
3	3	F	37		R	<1	No	Yes	Yes	Daily	ORIF of periprosthetic fracture, deep sepsis (<i>E. coli</i>), lost	Septic revision	2				Missing for interview but included
4	4	F	43	HIV, HCV	R	>1	Yes	Yes	Yes	No	Uneventful	Death	6				
5	5	M	31	HCV	R	<1	No	Yes	Yes	Occasionally	Uneventful	Death	73				
6	6	M	55	HCV	L	>10	No	Yes	Yes	Occasionally	Uneventful	Death	9				
7	7				R						Uneventful						
8	8	F	44		R	>10	No	No	No	Daily	Deep sepsis (CNS), died	Septic revision	1				
9	9	M	55		R	>1	No	No	Yes	No	Uneventful	Phone interview	98				Phone interview only
10	10	F	48	HCV	L	>1	No	No	Yes	No	Uneventful	Phone interview	24				
11	11	M	52	HCV	R	>1	No	Yes	Yes	Occasionally	Deep sepsis (SA), two-staged exchange	Septic revision	24				
12	12	F	28	HCV	L	<1	Yes	Yes	No	No	Deep infection (CNS, SA), resection arthroplasty	Septic revision	69				
13	13	M	46	HIV, HCV	R	>10	No	Yes	Yes	Occasionally	Uneventful	Clinical and radiological follow up	52	4	4	86	0
14	14	F	37		L	>1	No	Yes	No	No	Uneventful	Clinical and radiological follow up	24	5	8	71	1.6
15	15	M	41	HIV, HCV	R	>10	No	Yes	No	Occasionally	Uneventful	Clinical and radiological follow up	123	5	11	99	0
16	16	M	36	HIV	L	>1	No	No	Yes	Occasionally	Uneventful	Clinical and radiological follow up	113	5	3	99	0
17	17	M	32	HCV	L	<1	No	No	Yes	No	Uneventful	Clinical and radiological follow up	29	5	5	100	0.2

Table 2 continued

Patient	THA	Gender	Age	Positive serology (years)	Side	IIDU abstinence	IIDU recurrence	MCHPP	Smoking	Alcohol	Postoperative course	Endpoint	Time (months)	CRP	BSR	HHS	WOMAC
16	18	M	39		L	<1	Yes	No	Yes	Daily	Uneventful	Clinical and radiological follow up	80	5	9	100	0.4
17	19	F	39	HIV, HCV	L	>1	No	No	Yes	No	Uneventful	Clinical and radiological follow up	87	5	3	95	0.8
18	20	M	44	HIV, HCV	L	>1	No	Yes	Yes	No	Uneventful	Clinical and radiological follow up	86	15	38	96	1.5
19	21	M	48	HCV	R	>10	No	No	Yes	Daily	Uneventful	Clinical and radiological follow up	26	19	29	69	2
20	22	M	46	HCV	L	>10	No	No	Yes	Daily	<i>Stem revision due to periprosthetic fracture complicated with infection (CNS), debridement, one-stage exchange</i>	<i>Aseptic revision</i>	6	5	15	99	0
21	23	M	49	HIV, HCV	R	>1	No	Yes	Yes	Occasionally	<i>Stem revision due to aseptic loosening after unrecognized fracture</i>	<i>Aseptic revision</i>	11	5	21	100	0
22	24	M	50		R	>1	No	Yes	Yes	Daily	One-stage exchange due to deep infection (CNS)	Septic revision	1	16	25	89	0
23	25 26	M	44		L R	<1	Yes	Yes	Yes	No	Suspected deep infection	Awaiting septic revision	29	26	52	95	2
24	27	M	3	HCV	R	<1	Yes	Yes	Yes	Occasionally	Suspected deep infection	Awaiting septic revision	30				
											Deep infection with SA, recurrent infections after two-stage exchanges, resection arthroplasty	Septic revision	6	12	26	68	2

IIDU illicit injecting drug use, *MCHPP* medically controlled heroine prescription program, *HIV* human immunodeficiency virus, *HCV* hepatitis C virus, *CNS* coagulase negative *Staphylococcus*, *SA* *Staphylococcus aureus*, *ORIF* open reduction and internal fixation, *bold* patients with septic implant failure, *italics* patients with aseptic revision

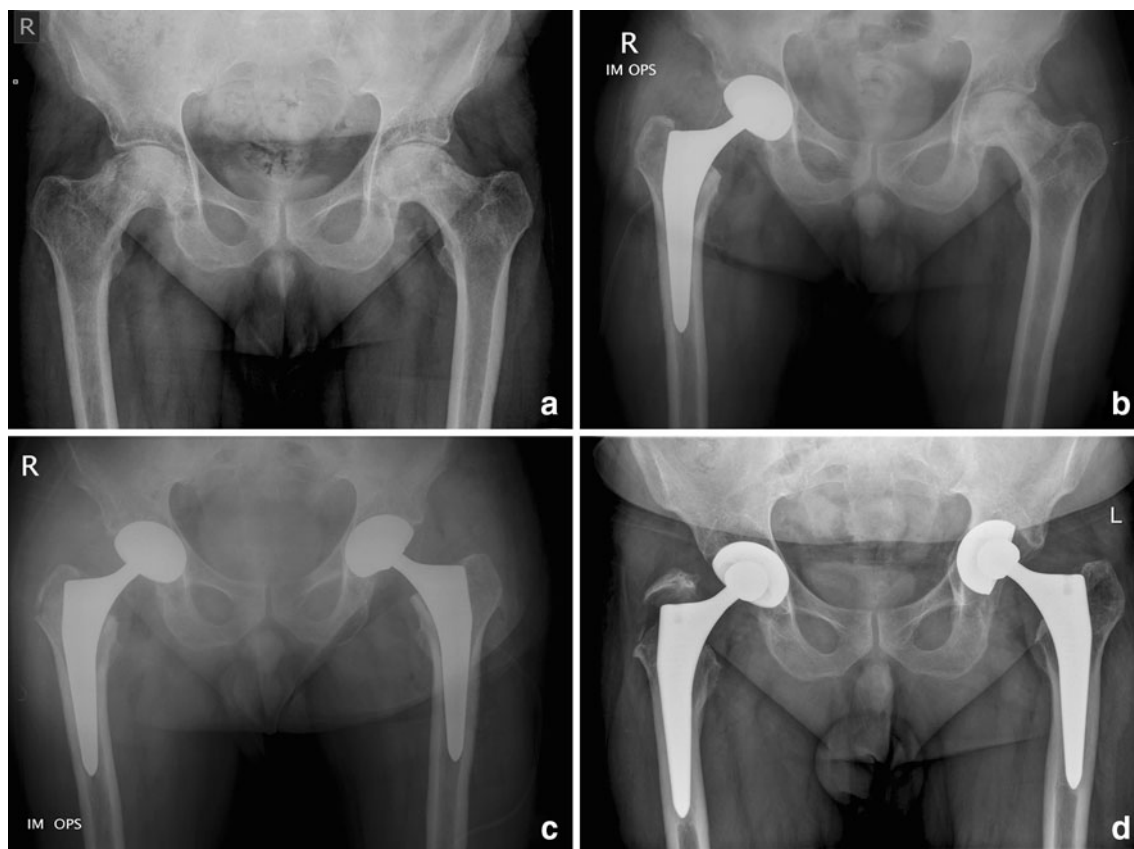


Fig. 1 X-ray of a 44-year-old man with bilateral osteonecrosis of the femoral head (a). A minimal invasive total hip arthroplasty was performed first on the right (b) and 6 weeks later on the left side (c). Postoperative X-rays showed a non-symptomatic avulsion fracture of the greater trochanter on the right side, which was treated

conservatively. 30 months after index procedure the patient presented elevated BSR and CRP levels as well as implant loosening, cup dislocation and gross metaphyseal and acetabular osteolysis on both sides (d). Although he denied joint aspiration and any kind of surgical treatment, both hips were considered as septic failures

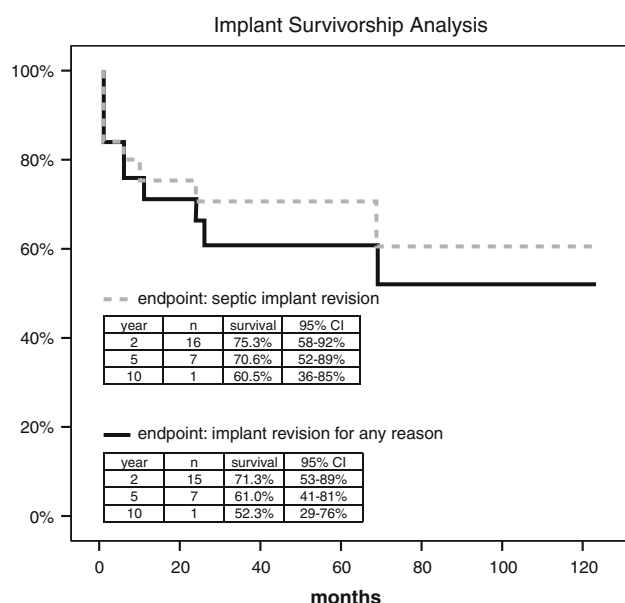


Fig. 2 Kaplan–Meier survivorship analysis with implant failure for septic and for any reasons as endpoints

According to our chart review, abstinence from illicit injection drug use at time of the index surgery was declared to be greater than 10 years in six, greater than 1 year in ten and less than 1 year in eight patients. Fourteen participated in a medically controlled heroine prescription program. The others were abstinent from illicit drug use without substitution. In 5 of 24 patients, recurrence of illicit injecting drug use was recorded either from the questionnaire or from the charts. All recurrences occurred in patients having stopped illicit drug use for less than 1 year before the index surgery ($p = 0.001$) and four of them participated in a medically controlled heroine prescription program ($p = 0.36$).

Four septic failures out of 5 THA's occurred in patients admitting illicit drug use recurrence and 3 septic failures out of 20 THA's in patients not admitting illicit drug use recurrence ($p = 0.023$). Five septic failures out of 8 THA's occurred in patients having stopped illicit drug use for less than 1 year and 3 out of 14 THA's in patients having stopped illicit drug use for more than 1 year ($p = 0.061$).

Seven patients had positive serology for HIV, 16 for HCV and 6 for both, all under a medically supervised treatment protocol. None of the THA's implanted in HIV-positive patients were revised because of deep sepsis.

At time of index surgery 19 patients were smoking at least ten cigarettes a day, 6 admitted to daily smoking and further 8 occasionally abused alcohol.

Discussion

In this retrospective analysis the 10-year cumulative septic implant revision rate of THA in patients with a history of illicit injecting drug use was as high as 39.5 % (CI: 15;64) and verified our concerns about an unacceptable high risk of septic complications in this population.

During the last decade, 1 % of primary THA's were performed in patients with a history of illicit injecting drug abuse in our institution. This is more than would be expected when considering the approximately 30,000 illicit injecting drug users in our country, representing about 0.4 % of our population. The urban location of our department and our practice to accept patients with a history of illicit injecting drug use for THA may explain this higher than expected rate.

With a mean age of 43 years, our patient population represents a relatively young group undergoing THA. This is most likely attributable to the etiology of osteoarthritis being femoral head osteonecrosis and septic arthritis in nearly half of the cases, which both are considered typical hip-specific complications of illicit injecting drug abuse and HIV.

Despite this young patient collective, four of 23 patients died during the follow-up period. Injecting drug use related deaths primarily occur among young adults and accounts for a large number of life years lost in developed societies. In Scotland and Spain for example, injecting drug related deaths accounts for 25–33 % of deaths of young adult males.

The high percentage of patients with positive serology for HIV (29 %) and HCV (54 %) is a known association in patients with illicit injecting drug use and can be explained by the sharing of syringes and other injection equipments. All HIV seropositive patients were under treatment of antiviral therapy and followed by the local department of infectious disease and none had symptoms of acquired immune deficiency syndrome.

Most disappointing was the 47.7 % implant failure rate at 10 years, 39.5 % due to deep sepsis. We are not aware of any other series that have reported the outcomes in a similar patient population; hence our results could be compared.

Looking to non-hemophilic HIV-positive patients, a higher rate of septic complications can yet not be

concluded from the literature [12–16]. However, THA in hemophilic HIV seropositive patients is indeed associated with a high septic revision rate, which in Hicks's series [12] was as high as 18.7 % in primary and 36.4 % in revision THA's. In one of these series [16] a subgroup of 5 HIV-positive illicit injecting drug users was included and 3 of them were complicated by deep sepsis after 1, 1.5 and 15 months. In our present series none of the patients sustaining deep sepsis was HIV seropositive and HIV therefore does not explain the high revision rate.

The risk to cause and spread infection with self-colonized strains has shown to be multiplied by injection or inhalation of drugs [8, 9]. Furthermore, the rate of periprosthetic joint infection after documented *Staphylococcus aureus* bacteremia is reported to range from 30 to 40 % [17]. It is, therefore, not surprising that injecting drug users more often sustain septic complications after THA's. In the present series, all patients affirmatively stated and their respective healthcare professionals confirmed abstinence from illicit injecting drug use at the time of the index procedure. Nevertheless, five patients admitted occasional recurrence of illicit injecting drug use after the index procedure. Abstinence of less than 1 year before index procedure was significantly associated with recurrence of illicit injecting drug use and both with septic failure of the prosthesis. However, if the patients with recurrence of illicit drug abuse were excluded, the septic failure rate would still reach 25 %. Although other risk factors for septic failure, such as younger age [18], alcohol abuse [19] and heavy smoking [20] were present in our group of patients, this still does not explain the exceedingly high complication rate.

From our data, we are unable to elucidate whether this high septic failure rate was due to the altered general health status of patients, who recently stopped injecting drug use thus making them more susceptible to infection or whether these patients are simply at higher risk of relapse, which they may not admit. In our study, drug tests [21, 22] in order to verify abstinence from illicit drug use were not performed. Additional limitations of the present study are its retrospective nature with difficulties of complete follow-up, the small group of patients with heterogenic indications for THA and different co-morbidities limiting statistical analysis. Furthermore, one can debate whether the origin of periprosthetic infection was hematogenous as opposed to an exogenous.

Despite these limitations valuable conclusions can be made: (1) THA's in patients with declared abstinence after a history of illicit injecting drug use results in an unacceptable high septic failure rate. (2) Declared abstinence of less than 1 year before THA is more often associated with recurrence of illicit drug abuse and both increase the risk of septic implant failure.

Therefore, from this study we have changed our current practice and have adopted the results as evidence when negotiating appropriate treatment of osteoarthritis with patients and their supporting health care professionals. We are also requesting hair analysis for drugs, documenting abstinence of at least 1 year before indicating THA.

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